

MACWPQLRLLLWKNLTFRRRQTCQLLLEVAWPLFIFLILISVRLSYPPYEQHECHFPNKA
(SEQ ID NO. 58). Given that there is an in-frame stop codon 6 to 9 nucleotides upstream from this location, the newly predicted start site is the first methionine codon that could produce a continuous open reading frame. Alignment of this new ABC1 cDNA sequence with related ABC transporter sequences ABCR and ABC-C (also known as ABC3) which also contain open reading frames for the 60 additional amino acids, indicates a high degree of similarity, implying that the homologous ABC transporter proteins begin with sequences related to the amino terminal extension sequence proposed for human ABC1. It is likely that the earlier published start site of the human ABC1 was predicted from the published mouse ABC1 cDNA sequence (Luciani et al., *Genomics*, 21:150-159 (1994); GenBank Accession no.: X75926) which contains an extra nucleotide "n" in the extension region such that the newly disclosed methionine is not in-frame. However, if the "n" nucleotide in the mouse sequence is ignored, the mouse and human sequences of the extension region are identical. In light of these results, it is likely that the full length human ABC1 protein contains 2261 amino acids rather than 2201 amino acids, as previously suggested by Langmann et al. and others. Accordingly, Langmann et al. do not present the full open reading frame of human ABC1."

On page 37, please replace the first paragraph (beginning with "sequences. For example,") of the originally-filed application with the following paragraph:

"sequences. For example, the full length 3' UTR (SEQ ID NO: 6) contains 46 sequences (AA)nCU/UC(AA)n (SEQ ID NO: 59) which have been shown to be necessary for binding of Vigilin. Vigilin, a ubiquitous protein with 14K homology domains, is the estrogen-inducible vitellogenin mRNA 3'-untranslated region binding protein (*J. Biol. Chem.*, 272: 12249-12252 (1997)). In addition to binding HDL, Vigilin has been shown to bind to the 3' flanking region of mRNAs and to increase the half-life of the mRNA transcript (*Mol. Cell. Biol.*, 18:3991-4003

B2 (1998)). Thus, the 3' flanking region could be altered, for example, to increase the binding of Vigilin, thereby increasing the half-life of the ABC1 mRNA. Preferably, the isolated polynucleotide comprises the sequence shown in SEQ ID NO: 4. Also preferably, the isolated polynucleotide comprises the sequence shown in SEQ ID NO: 5. In another preferred embodiment, the isolated polynucleotide comprises the sequence shown in SEQ ID NO: 6. In other preferred embodiments, the polynucleotide comprises a sequence that hybridizes, under stringent conditions, to the nucleotide sequence set forth in SEQ ID NO: 4, SEQ ID NO: 5, or SEQ ID NO: 6."

On page 59, please replace the first paragraph (beginning with "therapeutic effect") of the originally-filed application with the following paragraph:

B3 (K.F.) mm "therapeutic effect. A typical dosage may range from about 0.1 g/kg to about 100 mg/kg or more, depending on the factors mentioned above."

On page 76, please replace the third paragraph (beginning with "The amount of") of the originally-filed application with the following paragraph:

B4 (K.F.) mm "The amount of ABC1 protein can be assayed using any of the well-known methods of measuring protein. Preferably, the amount of ABC1 protein is measured using an immunoassay. In one embodiment, the amount of ABC1 protein is determined by (a) contacting the cell sample with a population of anti-ABC1 antibodies and (b) detecting the anti-ABC1 antibodies associated with the cell sample. For example, the ABC1 protein can be contacted with an antiserum raised against a synthetic peptide corresponding to KNQTVVDAVLTSFLQDEKVKES (SEQ ID NO. 60) located at the C-terminus, as described in Example 11. The anti-ABC1 antibodies can be detected using several methods known in the art, including, for example, western blotting, immunoprecipitation, and FACS, wherein the detection can be accomplished using radioactive,

B4
colorometric, or fluorescent labeling. One preferred method for measuring the amount of ABC1 protein in a cell sample is immunoprecipitation, wherein biotinylated ABC1 proteins are contacted with anti-ABC1 antibody and the bound anti-ABC1 antibody is detected using streptavidin horse radish peroxidase.”

On page 107, please replace the fourth paragraph (beginning with “To determine which”) of the originally-filed application with the following paragraph:

B5
“To determine which portion of the 5' flanking region of ABC1 retains transcriptional activity in response to nuclear ligands, various plasmids containing a different portion of the 5' flanking region and a luciferase reporter gene were transfected into RAW 264.7 cells treated with at least one ligand for the nuclear receptors. Using this system, an sterol response element corresponding to nucleotides 1480-1510 of SEQ ID NO: 3 was identified. The sterol response element contains a direct repeat-4 element TGACCGatagTAACCT (SEQ ID NO: 61). Confirmation of the sterol response element was obtained using site-directed mutagenesis and band-shift assay techniques.”

On page 108, please replace the third paragraph (beginning with “Site-Directed Mutagenesis”) of the originally-filed application with the following paragraph:

B6
“Site-Directed Mutagenesis: The sterol response element corresponding to nucleotides 1480-1510 of SEQ ID NO: 3 was mutated in the 1080-1643 sequence described above using site-directed mutagenesis. Specifically, the response element containing a direct repeat-4 element TGACCGatagTAACCT (SEQ ID NO: 61) was mutated to CTGCACatagTAACCT (SEQ ID NO: 62) using the GeneEditor system (Promega, Madison, WI) according to the manufacturer's protocol.”
